Attorney's Docket No.: 13425-0192US1 / BV-1087 US

Applicant: Giles Brown et al. Serial No.: 10/581,545 Filed: November 14, 2008

Page : 7 of 36

AMENDMENTS TO THE CLAIMS

This listing of claims replaces all prior versions and listings of claims in the application:

Listing of Claims:

1. (original) A method of synthesising a 2-substituted adenosine of formula I, which comprises converting 2-nitro-pentabenzoyl adenosine to the 2-substituted adenosine:

I

wherein $R = C_{1-6}$ alkoxy (straight or branched), a phenoxy group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_3 -, cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy), a benzyloxy group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_3 -, cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy), or a benzoyl group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_3 -, cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy).

- 2. (original) A method according to claim 1, wherein R = methoxy, ethoxy, propoxy, butoxy, pentyloxy, hexyloxy, phenoxy, benzyloxy, or benzoyl.
- (currently amended) A method according to claim 1, wherein 2-nitro-pentabenzoyl
 adenosine is converted to the 2-substituted adenosine by a process comprising reacting
 deprotection, and reaction with C₁₋₆ alkoxide anion, or a phenoxide anion and
 deprotecting protected functional groups.

Serial No.: 10/581,545 Filed: November 14, 2008

Page : 8 of 36

 (original) A method according to claim 3, wherein the anion is methoxide anion produced from MeOH/NaOMe, MeOH/n-BuLi, MeOH/NaOH, MeOH/NaH, or MeOH/KO^tBu.

- (previously presented) A method according to claim 1, which further comprises converting pentabenzoyl adenosine to 2-nitro-pentabenzoyl adenosine.
- 6. (original) A method of synthesising 2-nitro-pentabenzoyl adenosine which comprises converting pentabenzoyl adenosine to 2-nitro-pentabenzoyl adenosine.
- 7. (currently amended) A method according to claim 5, wherein pentabenzoyl adenosine is converted to 2-nitro-pentabenzoyl adenosine by nitrating pentabenzoyl adenosine by a process comprising reacting the pentabenzoyl adenosine with using tetrabutylammonium nitrate (TBAN), or tetramethylammonium nitrate (TMAN) as a nitrating reagent.
- (currently amended) A method according to claim 7, which further comprises reducing
 the amount of tetrabutylammonium nitrate TBAN or tetramethylammonium nitrate
 TMAN contaminating the 2-nitro-pentabenzoyl adenosine after the nitration reaction.
- (currently amended) A method according to claim 8, wherein <u>reducing</u> the amount of <u>tetrabutylammonium nitrate</u> TBAN or <u>tetramethylammonium nitrate</u> TMAN is reduced by is performed by a process comprising washing the 2-nitro-pentabenzoyl adenosine with water.
- (currently amended) A method according to claim 9, which further comprises recrystallising the 2-nitro-pentabenzoyl adenosine after the washing with water.
- 11. (previously presented) A method according to claim 5, which further comprises converting adenosine to pentabenzovl adenosine.
- 12. (original) A method of synthesising pentabenzoyl adenosine or 2-nitro-pentabenzoyl adenosine which comprises converting adenosine to pentabenzoyl adenosine.

Serial No.: 10/581,545 Filed: November 14, 2008

Page : 9 of 36

(currently amended) A method according claim 11, wherein the conversion of adenosine
to pentabenzoyl adenosine comprises reacting adenosine with benzoyl chloride is
benzoylated using benzoyl chloride.

14. (original) 2-nitro pentabenzoyl adenosine.

15-17. (cancelled)

- 18. (currently amended) A method of reducing the amount of <u>tetrabutylammonium nitrate</u>

 TBAN or <u>tetramethylammonium nitrate</u> TMAN contaminating 2-nitro-pentabenzoyl adenosine formed by nitration of pentabenzoyl adenosine with <u>tetrabutylammonium</u>

 <u>nitrate</u> TBAN or <u>tetramethylammonium nitrate</u> TMAN, which comprises washing the 2nitro-pentabenzoyl adenosine with water.
- (currently amended) A method according to claim 18 which further comprises
 recrystallising the 2-nitro-pentabenzoyl adenosine 2-ntiro-pentabenzoyl adenosine after
 washing with water.
- 20. (currently amended) A method of synthesising a 2-substituted adenosine of formula I, which comprises: nitrating adenosine pentaacetate using by a process comprising reacting adenosine pentaacetate with tetrabutylammonium nitrate (TBAN) or tetramethylammonium nitrate (TMAN) to produce 2-nitroadenosine pentaacetate; reducing the amount of tetrabutylammonium nitrate TBAN or tetramethylammonium nitrate TMAN contaminating the resulting 2-nitroadenosine pentaacetate; and then producing from the 2-nitroadenosine pentaacetate the 2-substituted adenosine from the 2-nitroadenosine pentaacetate to formula I:

Serial No.: 10/581,545 Filed: November 14, 2008

Page : 10 of 36

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wherein $R=C_{1-6}$ alkoxy (straight or branched), a phenoxy group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3} -, cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy), a benzyloxy group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3} -, cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy), or a benzoyl group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3} -, cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy).

- 21. (currently amended) A method according to claim 20, wherein the amount of tetrabutylammonium nitrate TBAN or tetramethylammonium nitrate TMAN contaminant is reduced by triturating the 2-nitroadenosine pentaacetate with isopropanol and washing the triturated 2-nitroadenosine pentaacetate with water.
- 22. (currently amended) A method according to claim 20, wherein the 2-substituted adenosine is produced from the 2-nitroadenosine pentaacetate by a process comprising deprotecting the 2-nitroadenosine pentaacetate and reaction reacting with a C₁₋₆ alkoxide anion or a phenoxide anion.
- 23. (currently amended) A method according to claim 20, wherein the 2-substituted adenosine is 2-methoxyadenosine 2-methoxyadenosine, and the 2-methoxyadenosine this is produced from the 2-nitroadenosine pentaacetate by reaction reacting the 2-nitroadenosine pentaacetate with methoxide anion from methanol/NaOMe, methanol/n-BuLi, methanol/NaOH, methanol/NaH, or methanol/KO¹Bu.
- 24. (previously presented) A method according to claim 20, which further comprises synthesising the adenosine pentaacetate by acylating adenosine.

Serial No.: 10/581,545 Filed: November 14, 2008

Page : 11 of 36

25. (currently amended) A method according to claim 24, wherein the adenosine is acylated to form an O-tri-acetyl and/or tetra-acetyl derivative of adenosine, the derivative(s) is isolated, and the isolated derivative(s) is <u>further</u> acylated to produce adenosine pentaacetate.

- 26. (previously presented) A method according to claim 24, which further comprises washing the adenosine pentaacetate to remove contaminating adenosine tetraacetate before nitrating the washed adenosine pentaacetate to form the 2-nitroadenosine pentaacetate.
- 27. (currently amended) A method of synthesising a 2-substituted adenosine of formula 1,

wherein $R = C_{1-6}$ alkoxy (straight or branched), a phenoxy group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3-} , cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy), a benzyloxy group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3-} , cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy), or a benzoyl group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3-} , cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy);

which wherein the method comprises acylating adenosine to form an O-tri-acetyl and/or tetra-acetyl derivative of adenosine, isolating the derivative(s), <u>further</u> acylating the isolated derivative(s) to produce adenosine pentaacetate, and producing the 2-substituted adenosine from the adenosine pentaacetate.

28. (currently amended) A method according to claim 27 which further comprises washing the adenosine pentaacetate to reduce the amount of contaminating adenosine tetraacetate

Attorney's Docket No.: 13425-0192US1 / BV-1087 US

Applicant: Giles Brown et al. Serial No.: 10/581,545 Filed: November 14, 2008

Page : 12 of 36

before producing the 2-substituted adenosine of formula I from the washed adenosine pentaacetate.

29. (currently amended) A method of synthesising a 2-substituted adenosine of formula I,

wherein $R = C_{1-6}$ alkoxy (straight or branched), a phenoxy group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3-} , cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy), a benzyloxy group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3-} , cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy), or a benzoyl group (unsubstituted, or mono-, or di-substituted by halo, amino, CF_{3-} , cyano, nitro, C_{1-6} alkyl, or C_{1-6} alkoxy);

which wherein the method comprises acylating adenosine, or an acylated derivative of adenosine, to form adenosine pentaacetate, washing the adenosine pentaacetate to reduce the amount of contaminating adenosine tetraacetate, and producing the 2-substituted adenosine of formula I from the washed adenosine pentaacetate.

- 30. (currently amended) A method according to claim 27, which further comprises nitrating the adenosine pentaacetate to produce 2-nitroadenosine pentaacetate, and producing the 2-substituted adenosine of formula I from the 2-nitroadenosine pentaacetate.
- 31. (currently amended) A method according to claim 30, wherein the 2-substituted adenosine is 2-methoxyadenosine, and the 2-methoxyadenosine is produced by reacting methoxide anion from methanol/NaOMe, methanol/n-BuLi, methanol/NaOH, methanol/NaH, or methanol/KO¹Bu with the 2-nitroadenosine pentaacetate.

Serial No.: 10/581,545 Filed: November 14, 2008

Page : 13 of 36

32. (previously presented) A method according to claim 20, which further comprises converting 2-nitroadenosine pentaacetate to 2-chloroadenosine pentaacetate before

producing the 2-substituted adenosine from the 2-chloroadenosine pentaacetate.

33. (original) A method of synthesising a 2-substituted adenosine, which comprises converting 2-chloroadenosine pentaacetate to the 2-substituted adenosine.

- 34. (original) A method according to claim 33, which further comprises producing the 2chloroadenosine pentaacetate from 2-nitroadenosine pentaacetate.
- 35. (previously presented) A method according to claim 32, wherein the 2-substituted adenosine is 2-methoxyadenosine, and the 2-chloroadenosine pentaacetate is converted to 2-methoxyadenosine by reaction with methoxide anion from methanol/NaOMe, methanol/n-BuLi, methanol/NaOH, or methanol/NaH with the 2-nitroadenosine pentaacetate.
- 36. (cancelled)
- (original) A method of synthesising 2-methoxyadenosine, which comprises reacting
 methoxide anion from methanol/NaOMe, methanol/n-BuLi, methanol/NaOH,
 methanol/NaH, or methanol/KO¹Bu with 2-nitroadenosine pentaacetate.
- 38. (currently amended) A method of synthesising 2-methoxyadenosine, which comprises the steps shown in scheme 1 or 2:

Serial No.: 10/581,545 Filed: November 14, 2008

Page : 14 of 36

Scheme 1

Attorney's Docket No.: 13425-0192US1 / BV-1087 US

Applicant: Giles Brown et al. Serial No.: 10/581,545 Filed: November 14, 2008

Page : 15 of 36

Adenosine

Adenosine pentaacetate

2-nitroadenosine pentaacetate

2-chloroadenosine pentaacetate

Scheme 2.

2-methoxyadenosine

(spongosine)

Serial No.: 10/581,545 Filed

: November 14, 2008

: 16 of 36 Page

40. (original) 2-methoxyadenosine which is >96% pure.

- 41. (currently amended) A method of synthesising 2-nitroadenosine pentaacetate, which comprises nitrating adenosine pentaacetate using by reaction with tetrabutylammonium nitrate TBAN or tetramethylammonium nitrate TMAN to produce 2-nitroadenosine pentaacetate, and reducing the amount of tetrabutylammonium nitrate TBAN or tetramethylammonium nitrate TMAN contaminating the 2-nitroadenosine pentaacetate.
- 42. (currently amended) A method according to claim 41, wherein the amount of tetrabutylammonium nitrate TBAN or tetramethylammonium nitrate TMAN contaminant is reduced by triturating the 2-nitroadenosine pentagetate with isopropanol and washing the triturated 2-nitroadenosine pentaacetate with water.
- 43. (currently amended) A method of synthesising adenosine pentaacetate, 2-nitroadenosine pentageetate, or a 2-substituted adenosine of formula I, which includes the comprising following steps: acylating adenosine to form an O-tri-acetyl and/or tetra-acetyl derivative of adenosine, isolating the derivative(s), and further acylating the isolated derivative(s) to produce adenosine pentaacetate.
- 44. (currently amended) A method of synthesising adenosine pentaacetate, 2-nitroadenosine pentaacetate, or a 2-substituted adenosine of formula I, which includes comprising the following steps: acylating adenosine or an acylated derivative of adenosine to form adenosine pentaacetate; and washing the adenosine pentaacetate to reduce the amount of contaminating adenosine tetraacetate.